

PATENT

IN THE UNITED STATES PATENT AND TRADEMARK OFFICE

Applicant: Dominic P. Behan : Art Unit: Not Yet Assigned  
Serial No.: Not Assigned : Examiner: Not Assigned  
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INFORMATION DISCLOSURE STATEMENT

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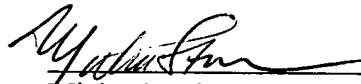
S I R :

Pursuant to 37 C.F.R. §§ 1.97 and 1.98 and to the duty of disclosure set forth in 37 C.F.R. § 1.56, the Examiner in charge of the above-identified application is requested to consider and make of record the references listed on the PTO/SB/08A submitted herewith.

Although the information submitted herewith may be "material" to the Examiner's consideration of the subject application, this submission is not intended to constitute an admission that such information is "prior art" as to the claimed invention.

In accordance with 37 C.F.R. § 1.97(g), the filing of this Information Disclosure Statement shall not be construed to mean that a search has been made.

Respectfully submitted,

  
\_\_\_\_\_  
Michael P. Straher  
Attorneys for Applicant

Enclosures:  
PTO/SB/08A

Dated: September 22, 2003

Cozen O'Connor  
1900 Market Street  
Philadelphia, PA 19103  
Tel: 215-665-2000

<b>Form PTO-1449 Modified</b>  List of Patent and Publications Cited by Applicant (Use several sheets if necessary)  U.S. Department of Commerce Patent and Trademark Office		Docket No. 5.US10.CON	Serial No. Not Assigned
		Applicant <b>Dominic P. Behan, et al.</b>	
		Filing Date Herewith	Group Not Yet Assigned
<b>OTHER DOCUMENTS (Including Author, Title, Date, Pertinent Pages, Etc.)</b>			
	AA	Gdaniec, M., et al. "Crystal and molecular structure of 3-(4-chlor=benzoylimino)-5,6-dihydro-3H-imidazo[2,1-c]-1,2,4-dithiazole," <i>Chemical Abstracts</i> XP-002177674, 1986, 111(20), 1 page	
	AB	Gdaniec, M., et al., "Crystal and molecular structure of 3-(4-chlorobenzoylimino)-5,6-dihydro-3H-imidazo[2,1-c]-1,2,4-dithiazole," <i>Chemical Abstracts</i> XP-002177676, 1989, 19(3), 1 page	
	AC	Saczewski, F., "Synthesis, transformations and tuberculostatic activity of 1-(N-arylthiocarbamoyl)imidazolidine-2-thiones," <i>Chemical Abstracts</i> XP-002177673, 1993, 121(7), 1 page	
	AD	Saczewski, F., "Synthesis transformations and tuberculostatic activity of 1-arylthiocarbamoyl)imidazolidine-2-thiones," <i>Chemical Abstracts</i> XP-002177675, 1993, 1 page	
	AE	Saczewski, F., et al., "Synthesis and transformations of 3-benzoylimino-5,6-dihydro-3H-imidazo[2,1-c][1,2,4]dithiazole," <i>Communications</i> , September 1986, 751-753	
	AF	PCT International Search Report dated October 9, 2001	
EXAMINER		DATE CONSIDERED	

<b>Form PTO-1449 Modified</b>  List of Patent and Publications Cited by Applicant (Use several sheets if necessary)  U.S. Department of Commerce Patent and Trademark Office				Docket No. 5.0510.CON		Serial No. Not Assigned	
				Applicant Dominic P. Behan, et al.			
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<b>U. S. PATENT DOCUMENTS</b>							
<b>Examiner Initial</b>		<b>Document No.</b>	<b>Date</b>	<b>Name</b>	<b>Class</b>	<b>Subclass</b>	
<b>FOREIGN PATENT DOCUMENTS</b>							
<b>Examiner Initial</b>		<b>Document No.</b>	<b>Date</b>	<b>Country</b>	<b>Translation</b> YES NO		
	AG	WO 99/09024	02/25/99	PCT			
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		Applicant Dominic P. Behan et al.	
		Filing Date Herewith	Group Not Yet Assigned
<b>OTHER DOCUMENTS (Including Author, Title, Date, Pertinent Pages, Etc.)</b>			
	BA	Bertin, B. et al., "Cellular signaling by an agonist-activated receptor/Gs-alpha fusion protein", <i>Proc. Natl. Acad. Sci. USA</i> , 1994, 91, 8827-8831.	
	BB	Seifert, R. et al., "Different effects of Gs alpha splice variants on beta-2-adrenoreceptor-mediated signaling, <i>J. Biol. Chem.</i> , 1998, 273, 5109-5116.	
	BC	Wise, A. et al., "Rescue of functional interactions between the alpha-2A-adrenoreceptor and acylation-resistant forms of Gil-alpha by expressing the proteins from chimeric open reading frames," <i>J. Biol. Chem.</i> , 1997, 272, 24673-24678.	
	BD	Burt, A.R., et al., "Agonist occupation of an alpha-2A-adrenoreceptor-Gil-alpha fusion protein results in activation of both receptor-linked and endogenous Gi proteins", <i>J. Biol. Chem.</i> , 1997, 273, 10367-10375.	
<b>EXAMINER</b>		<b>DATE CONSIDERED</b>	

<b>Form PTO-1449 Modified</b>  List of Patent and Publications Cited by Applicant (Use several sheets if necessary)  U.S. Department of Commerce Patent and Trademark Office				Docket No. 57US10.2CON		Serial No. Not Assigned	
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<b>Examiner Initial</b>		<b>Document No.</b>	<b>Date</b>	<b>Country</b>	<b>Translation YES NO</b>		
	BE	WO 98/46995	22.10.98	PCT	X		
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	CA	2,135,253	08.05.96	Canada	X		
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<b>OTHER DOCUMENTS (Including Author, Title, Date, Pertinent Pages, Etc.)</b>			
	DA	Alla, S.A. et al., "Extracellular domains of the bradykinin B2 receptor involved in ligand binding and agonist sensing defined by anti-peptide antibodies," <i>J. Biol. Chem.</i> , 1996, 271, 1748-1755	
	DB	Advenier, C. et al., "Effects on the isolated human bronchus of SR 48968, a potent and selective nonpeptide antagonist of the neurokinin A (NK <sub>2</sub> ) receptors," <i>Am. Rev. Respir. Dis.</i> , 1992, 146(5, Pt. 1), 1177-1181	
	DC	Alexander, W.S. et al., "Point mutations within the dimer interfact homology domain of c-Mpl induce constitutive receptor activity and tumorigenicity," <i>EMBO J.</i> , 1995, 14(22), 5569-5578	
	DD	Arvanitakis, L. et al., "Human herpesvirus KSHV encodes a constitutively active G-protein-coupled receptor linked to cell proliferation," <i>Nature</i> , 1997, 385, 347-349	
	DE	Barker, E.L. et al., "Constitutively active 5-hydroxytryptamine <sub>2C</sub> receptors reveal novel inverse agonist activity of receptor ligands," <i>J. Biol. Chem.</i> , 1994, 269(16), 11687-11690	
	DF	Baxter, G., "5-HT <sub>2</sub> receptors: a family re-united?" <i>Trends Pharmacol. Sci.</i> , 1995, 16, 105-110	
	DG	Besmer, P. et al., "A new acute transforming feline retrovirus and relationship of its oncogene v-kit with the protein kinase gene family," <i>Nature</i> , 1986, 320, 415-421	
	GH	Blin, N. et al., "Mapping of single amino acid residues required for selective activation of G <sub>q/11</sub> by the m3 muscarinic acetylcholine receptor," <i>J. Biol. Chem.</i> , 1995, 270, 17741-17748	
	AI DI	Bond, R.A. et al., "Inverse agonists and G-protein-coupled receptors," in <i>Receptor-Based Drug Design</i> , Leff, P. (ed.), New York, M. Dekker, 1998, 363-377	
	DJ	Boone, C. et al., "Mutations that alter the third cytoplasmic loop of the a-factor receptor lead to a constitutive and hypersensitive phenotype," <i>Proc. Natl. Acad. Sci. USA</i> , 1993, 90(21), 9921-9925	
EXAMINER		DATE CONSIDERED	



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	DK	Burstein, E.S. et al., "Constitutive activation of chimeric m2/m5 muscarinic receptors and delineation of G-protein coupling selectivity domains," <i>Biochem. Pharmacol.</i> , 1996, 51(4), 539-544	
	DL	Burstein, E.S. et al., "Amino acid side chains that define muscarinic receptor/G-protein coupling. Studies of the third intracellular loop," <i>J. Biol. Chem.</i> , 1996, 271(6), 2882-2885	
	DM	Burstein, E.S. et al., "Constitutive activation of muscarinic receptors by the G-protein G <sub>q</sub> ," <i>FEBS Lett.</i> , 1995, 363(3), 261-263	
	DN	Bylund, D., "International union of pharmacology nomenclature of adrenoceptors," <i>Pharmacol. Rev.</i> , 1994, 46, 121-136	
	DO	Casey, C. et al., "Constitutively active mutant 5-HT <sub>2A</sub> serotonin receptors: inverse agonist activity of classical 5HT <sub>2A</sub> antagonists," <i>Soc. Neurosci.</i> , 1996, Abstract #699.10	
	DP	Cheatham, B. et al., "Substitution of the <i>erbB-2</i> oncoprotein transmembrane domain activates the insulin receptor and modulates the action of insulin-receptor substrate 1," <i>Proc. Natl. Acad. Sci. USA</i> , 1993, 90, 7336-7340	
	DQ	Chen, J. et al., "Tethered Ligand Library for Discovery of Peptide Agonists," <i>J. Biol. Chem.</i> , 1995, 270, 23398-23401	
	DR	Chen, T.S. et al., "Microbial hydroxylation and glucuronidation of the angiotensin II (AII) receptor antagonist MK 954," <i>J. Antibiot. (Tokyo)</i> , 1993, 46(1), 131-134	
	DS	Chen, W. et al., "A colorimetric assay for measuring activation of G <sub>s</sub> - and G <sub>q</sub> -coupled signaling pathways," <i>Anal. Biochem.</i> , 1995, 226(2), 349-354	
	DT	Chidiac, P. et al., "Inverse agonist activity of $\beta$ -adrenergic antagonists," <i>J. Pharm. Exp. Ther.</i> , 1994, 45, 490-499	
	DU	Clozel, M. et al., "In vivo pharmacology of Ro 46-2005, the first synthetic nonpeptide endothelin receptor antagonist: implications for endothelin physiology," <i>J. Cardiovas. Pharmacol.</i> , 1993, 22(Suppl. 8), S377-S379	
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	DV	Collesi, C. et al., "A splicing variant of the <i>RON</i> transcript induces constitutive tyrosine kinase activity and an invasive phenotype," <i>Mol. Cell. Biol.</i> , 1996, 16(2), 5518-5526	
	DW	Cooper, C.S. et al., "Molecular cloning of a new transforming gene from a chemically transformed human cell line," <i>Nature</i> , 1984, 311, 29-33	
	DX	De Dios, I. et al., "Effect of L-364,718 (CCK Receptor Antagonist) on Exocrine Pancreatic Secretion of Hydrocortison-Treated Rats," <i>Pancreas</i> , 1994, 9(2), 212-218	
	DY	Desbios-Mouthon, C. et al., "Deletion of Asn <sup>281</sup> in the $\alpha$ -subunit of the human insulin receptor causes constitutive activation of the receptor and insulin desensitization," <i>J. Clin. Endocrinol. Metab.</i> , 1996, 81(2), 719-727	
	DZ	Di Renzo, M.F. et al., "Expression of the Met/HGF receptor in normal and neoplastic human tissues," <i>Oncogene</i> , 1991, 6(11), 1997-2003	
	DAA	Di Renzo, M.F. et al., "Overexpression of the c-MET/HGF receptor gene in human thyroid carcinomas," <i>Oncogene</i> , 1992, 7, 2549-2553	
	DAB	Duprez, L. et al., "Germline mutations of the thyrotropin receptor gene cause non-autoimmune autosomal dominant hyperthyroidism," <i>Nature Genetics</i> , 1994, 7, 396-401	
	DAC	Eggericksx, D. et al., "Molecular Cloning of an Orphan G-Protein-Coupled Receptor that Constitutively Activates Adenylate Cyclase," <i>Biochem. J.</i> , 1995, 309, 837-843	
	DAE	Evans, B.E. et al., "Orally Active, Nonpeptide Oxytocin Antagonists," <i>J. Med. Chem.</i> , 1992, 35, 3919-3927	
	DAF	Fu, M. et al., "Functional autoimmune epitope on $\alpha_1$ -adrenergic receptors in patients with malignant hypertension," <i>Lancet</i> , 1994, 344, 1660-1663	
	DAG	Furitsu, T. et al., "Identification of Mutations in the Coding Sequence of the Proto-oncogene <i>c-kit</i> in a Human Mast Cell Leukemia Cell Line Causing Ligand-independent Activation of <i>c-kit</i> Product," <i>J. Clin. Invest.</i> , 1993, 92, 1736-1744	
	DAH	Gellai, M. et al., "Nonpeptide Endothelin Receptor Antagonists V: Prevention and Reversal of Acute Renal Failure in the Rat by SB 209670," <i>J. Pharm. Exp. Therap.</i> , 1995, 275(1), 200-206	
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<b>OTHER DOCUMENTS (Including Author, Title, Date, Pertinent Pages, Etc.)</b>			
	DAI	Gitter, B. et al., "Pharmacological Characterization of LY303870: A Novel Potent and Selective Nonpeptide Substance P (Neurokinin-1) Receptor Antagonist," <i>J. Pharm. Exp. Therp.</i> , 1995, 275(2), 737-744	
	DAJ	Gouilleux-Gruart, V. et al., "STAT-Related Transcription Factors are Constitutively Activated in Peripheral Blood Cells from Acute Leukemia Patients," <i>Blood</i> , 1996, 87(5), 1692-1697	
	DAK	Hansson, J.H. et al., "Hypertension caused by a truncated epithelial sodium channel $\gamma$ subunit: genetic heterogeneity of Liddle syndrome," <i>Nat. Genet.</i> , 1995, 11(1), 76-82	
	DAL	Hasegawa, H. et al., "Two Isoforms of the Prostaglandin E Receptor EP3 Subtype Different in Agonist-independent Constitutive Activity," <i>J. Biol. Chem.</i> , 1996, 271(4), 1857-1860	
	DAM	Hendler, F. et al., "Human Squamous Cell Lung Cancers Express Increased Epidermal Growth Factor Receptors," <i>J. Clin. Invest.</i> , 1984, 74, 647-651	
	DAN	Herrick-Davis, K. et al., "Constitutively Active 5HT <sub>2C</sub> Serotonin Receptor Created by Site-Directed Mutagenesis," <i>Soc. Neurosci.</i> , Abstract No. 699.18	
	DAO	Hieble, J., "International union of pharmacology. X. Recommendation for nomenclature of 1-adrenoceptors," <i>Pharm. Rev.</i> , 1995, 47, 267-270	
	DAP	Hill, S., "Distribution, Properties, and Functional Characteristics of Three Classes of Histamine Receptor," <i>Am. Soc. Pharm. Exp. Therap.</i> , 1990, 42(1), 45-83	
	DAQ	Högger, P. et al., "Activating and Inactivating Mutations in - and C-terminal i3 Loop Junctions of Muscarinic Acetylcholine Hm1 Receptors," <i>J. Biol. Chem.</i> , 1995, 270(13), 7405-7410	
	DAR	Ikeda, H. et al., "Expression and Functional Role of the Proto-oncogene c-kit in Acute Myeloblastic Leukemia Cells," <i>Blood</i> , 1991, 78(11), 2962-2968	
	DAS	Imura, R. et al., "Inhibition by HS-142-1, a novel nonpeptide atrial natriuretic peptide antagonist of microbial origin, of atrial natriuretic peptide-induced relaxation of isolated rabbit aorta through the blockade of guanylyl cyclase-linked receptors," <i>Mol. Pharm.</i> , 1992, 42, 982-990	
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	DAT	Jakubik, J. et al., "Constitutive activity of the M <sub>1</sub> -M <sub>4</sub> subtypes of muscarinic receptors in transfected CHO cells and of muscarinic receptors in the heart cells revealed by negative antagonists," <i>FEBS Letts.</i> , 1995, 377, 275-279	
	DAU	Kjelsberg, M.A. et al., "Constitutive activation of the $\alpha_{1B}$ -adrenergic receptor by all amino acid substitutions at a single site," <i>J. Biol. Chem.</i> , 1992, 267(3), 1430-1433	
	DAV	Knapp, R. et al., "Molecular biology and pharmacology of cloned opioid receptors," <i>FASEB J.</i> , 1995, 9, 516-525	
	DAW	Kosugi, S. et al., "Characterization of heterogeneous mutations causing constitutive activation of the luteinizing hormone receptor in familial male precocious puberty," <i>Human Mol. Genetics</i> , 1995, 4(2), 183-188	
	DAX	Kosugi, S. et al., "Identification of Thyroid-Stimulating Antibody-Specific Interaction Sites in the N-Terminal Region of the Thyrotropin Receptor," <i>Mol. Endocrinology</i> , 1993, 7, 114-130	
	DAY	Kraus, M. et al., "Demonstration of ligand-dependent signaling by the <i>erbB-3</i> tyrosine kinase and its constitutive activation in human breast tumor cells," <i>Proc. Natl. Acad. Sci. USA</i> , 1993, 90, 2900-2904	
	DAZ	Kudlacz, E. et al., "In Vitro and In Vivo Characterization of MDL 105,212A, a Nonpeptide NK-1/NK-2 Tachykinin Receptor Antagonist," <i>J. Pharm. Exp. Therap.</i> , 1996, 277(2), 840-851	
	DBA	Kuriu, A. et al., "Proliferation of Human Myeloid Leukemia Cell Line Associated with the Tyrosine-Phosphorylation and Activation of the Proto-oncogene <i>c-kit</i> Product," <i>Blood</i> , 1991, 78(11), 2834-2840	
	DBB	Labbe-Jullie, C. et al., "Effect of the nonpeptide neurotensin antagonist, SR 48692, and two enantiomeric analogs, SR 48527 and SR 49711, on neurotension binding and contractile responses in guinea pig ileum and colon," <i>J. Pharm. Exp. Therap.</i> , 1994, 271(1), 267-276	
	DBC	Latronico, A. et al., "A novel mutation of the luteinizing hormone receptor gene causing male gonadotropin-independent precocious puberty," <i>J. Clin. Endocrinol. Metabl.</i> , 1995, 80(8), 2490-2494	
<b>EXAMINER</b>		<b>DATE CONSIDERED</b>	

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	DBD	Laue, L. et al., "Genetic heterogeneity of constitutively activating mutations of the human luteinizing hormone receptor in familial male-limited precocious puberty," <i>Proc. Natl. Acad. Sci USA</i> , 1995, 92, 1906-1910	
	DBE	Løvlie, R. et al., "The Ca <sup>2+</sup> -sensing receptor gene (PCAR1) mutation T151M in isolated autosomal dominant hypoparathyroidism," <i>Hum. Genet</i> , 1996, 98, 129-133	
	DBF	Lefkowitz, R. et al., "Constitutive activity of receptors coupled to guanine nucleotide regulatory proteins," <i>Trends Pharmacol. Sci.</i> , 1993, 14, 303-307	
	DBG	Libermann, T. et al., "Amplification, enhanced expression and possible rearrangement of EGF receptor gene in primary human brain tumours of glial origin," <i>Nature</i> , 1985, 313, 144-147	
	DBH	Liu, C. et al., "Overexpression of c-met proto-oncogene but not epidermal growth factor receptor or c-erbB-2 in primary human colorectal carcinomas," <i>Oncogene</i> , 1992, 7, 181-185	
	DBI	Liu, J. et al., "Molecular mechanisms involved in muscarinic acetylcholine receptor-mediated G protein activation studied by insertion mutagenesis," <i>J. Biol. Chem.</i> , 1996, 271(11), 6172-6178	
	DBJ	Lonardo, F. et al., "The normal erbB-2 product is an atypical receptor-like tyrosine kinase with constitutive activity in the absence of ligand," <i>New Biologist</i> , 1990, 2(11), 992-1003	
	DBK	Maenhaut, C. et al., "RDC8 codes for an adenosine A2 receptor with physiological constitutive activity," <i>Biochem. Biophys. Res. Comm.</i> , 1990, 173(3), 1169-1178	
	DBL	Mann, J. et al., "Increased serotonin <sub>2</sub> and β-adrenergic receptor binding in the frontal cortices of suicide victims," <i>Arch. Gen. Psychiatry</i> , 1986, 43, 954-959	
	DBM	Martone, R.L. et al., "Human CRF receptor chimeras: Mapping of ligand binding determinants," 26th Meeting of the Society of Neuroscience, Washington, D.C. November 16-21, 1996, Abstract No. 609.8	
	DBN	Magnusson, Y. et al., "Autoimmunity in idiopathic dilated cardiomyopathy," <i>Circulation</i> , 1994, 89, 2760-2767	
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	DBO	Matus-Leibovitch, N. et al., "Truncation of the thyrotropin-releasing hormone receptor carboxyl tail causes constitutive activity and leads to impaired responsiveness in <i>Xenopus</i> Oocytes and AtT20 Cells," <i>J. Biol. Chem.</i> , 1995, 270(3), 1041-1047	
	DBP	Myles, G.M. et al., "Tyrosine 569 in the c-Fms juxtamembrane domain is essential for kinase activity and macrophage colony-stimulating factor-dependent internalization," <i>Mol. Cell. Biol.</i> , 1994, 14(7), 4843-4854	
	DBQ	Nanevicz, T. et al., "Thrombin receptor activating mutations," <i>J. Biol. Chem.</i> , 1996, 271(2), 702-706	
	DBR	Natali, P.G. et al., "Expression of the c-Met/HGF receptor in human melanocytic neoplasms: demonstration of the relationship to malignant melanoma tumour progression," <i>Br. J. Cancer</i> , 1993, 68, 746-749	
	DBS	Neilson, K.M. et al., "Constitutive activation of fibroblast growth factor receptor-2 by a point mutation associated with Crouzon syndrome," <i>J. Biol. Chem.</i> , 1995, 270(44), 26037-26040	
	DBT	Oda, S. et al., "Pharmacological profile of HS-142-1, a novel nonpeptide atrial natriuretic peptide (ANP) antagonist of microbial origin. II. Restoration by HS-142-1 of ANP-induced inhibition of aldosterone production in adrenal glomerulosa cells," <i>J. Pharm. Exp. Ther.</i> , 1992, 263(1), 241-245	
	DBU	O'Dowd, B.F. et al., "Site-directed mutagenesis of the cytoplasmic domains of the human $\beta_2$ -adrenergic receptor," <i>J. Biol. Chem.</i> , 1988, 263(31), 15985-15992	
	DBV	Offermanns, S. et al., " $G_{\alpha_{15}}$ and $G_{\alpha_{16}}$ Couple a Wide Variety of Receptors to Phospholipase C," <i>J. Biol. Chem.</i> , 1995, 270, 15175-15180	
	DBW	Palkowitz, A.D. et al., "Structural evolution and pharmacology of a novel series of triacid angiotensin II receptor antagonists," <i>J. Med. Chem.</i> , 1994, 37, 4508-4521	
	DBX	Parent, J. et al., "Mutations of two adjacent amino acids generate inactive and constitutively active forms of the human platelet-activating factor receptor," <i>J. Biol. Chem.</i> , 1996, 271(14), 7949-7955	
	DBY	Parfitt, A.M. et al., "Hypercalcemia due to constitutive activity of the parathyroid hormone (PTH)/PTH-related peptide receptor: comparison with primary hyperparathyroidism," <i>J. Clin. Endocr. Metab.</i> , 1996, 81, 3584-3588	
<b>EXAMINER</b>		<b>DATE CONSIDERED</b>	

<b>Form PTO-1449 Modified</b>  List of Patent and Publications Cited by Applicant (Use several sheets if necessary)  U.S. Department of Commerce Patent and Trademark Office		Docket No. 5.US10.CON	Serial No. Not Assigned
		Applicant Dominic P. Behan et al.	
		Filing Date Herewith	Group Not Yet Assigned
<b>OTHER DOCUMENTS (Including Author, Title, Date, Pertinent Pages, Etc.)</b>			
	DBZ	Samama, P. et al., "A mutation-induced activation state of the $\beta$ 2-adrenergic receptor," <i>J. Biol. Chem.</i> , 1993, 268(7), 4625-4636	
	DCA	Sautel, M. et al., "Neuropeptide Y and the nonpeptide antagonist BIBP 3226 share an overlapping binding site at the human Y1 receptor," <i>Am. Soc. Pharm. Exp. Ther.</i> , 1996, 50, 285-292	
	DCB	Sawutz, D.G. et al., "Pharmacology and structure-activity relationships of the nonpeptide bradykinin receptor antagonist WIN 64338," <i>Can. J. Physiol. Pharmacol.</i> , 1995, 73, 805-811	
	DCC	Scheer, A. et al., "Constitutively active G protein-coupled receptors: potential mechanisms of receptor activation," <i>J. Rec. Signal Transduct. Res.</i> , 1997, 17(1-3), 57-73	
	DCD	Scheer, A. et al., "The activation process of the $\alpha_{1B}$ -adrenergic receptor: Potential role of protonation and hydrophobicity of a highly conserved aspartate," <i>Proc. Natl. Acad. Sci. USA</i> , 1997, 94, 808-813	
	DCE	Schwinn, D.A. et al., "Cloning and pharmacological characterization of human <i>Alpha</i> -1 adrenergic receptors: sequence corrections and direct comparison with other species homologues," <i>J. Pharmacol.</i> , 1995, 272(1), 134-142	
	DCF	Schild, L. et al., "A mutation in the epithelial sodium channel causing Liddle disease increases channel activity in the <i>Xenopus laevis</i> oocyte expression system," <i>Proc. Natl. Acad. Sci. USA</i> , 1995, 92, 5699-5703	
	DCG	Seeman, P. et al., "Dopamine receptor pharmacology," <i>Trends Pharmacol. Sci.</i> , 1994, 15, 264-270	
	DCH	Seeman, P. et al., "Dopamine D4 receptors elevated in schizophrenia," <i>Nature</i> , 1993, 365, 441-445	
	DCI	Serradeil-Le Gale, C. et al., "Biochemical and pharmacological properties of SR 49059, a new, potent, nonpeptide antagonist of rat and human vasopressin $V_{1a}$ receptors," <i>J. Clin. Invest.</i> , 1993, 92, 224-231	
	DCJ	Sharif, M. et al., "Malignant transformation by G protein-coupled hormone receptors," <i>Mol. Cell. Endocrinology</i> , 1994, 100, 115-119	
EXAMINER		DATE CONSIDERED	

<b>Form PTO-1449 Modified</b>  List of Patent and Publications Cited by Applicant (Use several sheets if necessary)  U.S. Department of Commerce Patent and Trademark Office		Docket No. 5-8510-CON	Serial No. Not Assigned
		Applicant Dominic P. Behan et al.	
		Filing Date Herewith	Group Not Yet Assigned
<b>OTHER DOCUMENTS (Including Author, Title, Date, Pertinent Pages, Etc.)</b>			
	DCK	Jensen et al., "mRNA Profiling of Rat Islet Tumors Reveals Nkx 6.1 as a $\beta$ -Cell-specific Homeodomain Transcription Factor," <i>J. Biol. Chem.</i> , 1996, 271(31), 18749-18758	
	DCL	Konig et al., "Method for Identifying Ligands That Bind to Cloned G <sub>s</sub> - or G <sub>i</sub> -Coupled Receptors," <i>Mol. Cell. Neuro.</i> , 1991, 2, 331-337	
	DCM	Leonard, J. et al., "The LIM family transcription factor Isl-1 requires cAMP response element binding protein to promote somatostatin expression in pancreatic islet cells," <i>Proc. Natl. Acad. Sci. USA</i> , 1992, 89, 6247-6251	
	DCN	Marchese, A. et al., "Cloning of Human Genes Encoding Novel G Protein-Coupled Receptors," <i>Genomics</i> , 1994, 23, 609-618	
	DCO	Marks, D.L. et al., "Simultaneous Visualization of Two Cellular mRNA Species in Individual Neurons by Use of a New Double <i>in Situ</i> Hybridization Method," <i>Mol. &amp; Cell. Neuro.</i> , 1992, 3, 395-405	
	DCP	O'Dowd, B. et al., "Cloning and chromosomal mapping of four putative novel human G-protein-coupled receptor genes," <i>Gene</i> , 1997, 187, 75-81	
	DCQ	Sakurai T. et al., "Orexins and Orexin Receptors: A Family of Hypothalamic Neuropeptides and G Protein-Coupled Receptors that Regulate Feeding Behavior," <i>Cell</i> , 1998, 92, 573-585	
	DCR	Song, Z.-H. et al., "Molecular Cloning and Chromosomal Localization of Human Genes Encoding Three Closely Related G Protein-Coupled Receptors," <i>Genomics</i> , 1995, 28, 347-349	
	DCS	Suzuki, M. et al., "Regulatable Promoters for Use in Gene Therapy Applications: Modification of the 5'-Flanking Region of the CFTR Gene with Multiple cAMP Response Elements to Support Basal, Low-Level Gene Expression that can be Upregulated by Exogenous Agents that Raise Intracellular Levels of cAMP," <i>Human Gene Therapy</i> , 1996, 7, 1883-1893	
	DCT	Xu, Y. et al., "Identification of Human OGR1, a Novel G Protein-Coupled Receptor That Maps to Chromosome 14," <i>Genomics</i> , 1996, 35, 397-402	
*	DCU	Nichols, J.G. et al. (eds.), "Indirect Mechanisms of Synaptic Transmission," in <i>From Neuron To Brain</i> , 3rd Edition, Sinauer Associates, Inc., 1992	
	DCV	Osl et al. (eds.), in <i>Remington's Pharmaceutical Sciences</i> , 16th Edition, Mack Publishing Co., 1980	
EXAMINER		DATE CONSIDERED	

A copy of this reference will not be forwarded to the U.S. Patent and Trademark Office since it is believed to be too voluminous and easily obtainable by the Examiner.



<b>Form PTO-1449 Modified</b>  List of Patent and Publications Cited by Applicant (Use several sheets if necessary)  U.S. Department of Commerce Patent and Trademark Office		Docket No. 5. US10,700	Serial No. Not Assigned
		Applicant Dominic P. Behan et al.	
		Filing Date Herewith	Group Not Yet Assigned
<b>OTHER DOCUMENTS (Including Author, Title, Date, Pertinent Pages, Etc.)</b>			
	DCW	Parma, J. et al., "Somatic mutations in the thyrotropin receptor gene cause hyperfunctioning thyroid adenomas," <i>Nature</i> , 1993, 365, 649-651	
	DCX	Pei, G. et al., "A constitutive active mutant $\beta_2$ -adrenergic receptor is constitutively desensitized and phosphorylated," <i>Proc. Natl. Acad. Sci. USA</i> , 1994, 91, 2699-2702	
	DCY	Pendley, C.E. et al., "The gastrin/cholecystokinin-B receptor antagonist L-365,260 reduces basal acid secretion and prevents gastrointestinal damage induced by aspirin, ethanol and cysteamine in the rat," <i>J. Pharmacol. Exp. Ther.</i> , 1993, 265(3), 1348-1354	
	DCZ	Peroutka, S., "Serotonin receptor subtypes. Their evolution and clinical relevance," <i>CNS Drugs</i> , 1995, 4 (Suppl. 1), 18-27	
	DDA	Pettibone, D.J. et al., "Development and pharmacological assessment of novel peptide and nonpeptide oxytocin antagonists," <i>Regul. Pept.</i> , 1993, 45, 289-293	
	ddb	Prat, M.P. et al., "The receptor encoded by the human c-Met oncogene is expressed in hepatocytes, epithelial cells and solid tumors," <i>Int. J. Cancer</i> , 1991, 49, 323-328	
	DDC	Prezeua, L. et al., "Changes in the carboxy-terminal domain of metabotropic glutamate receptor 1 by alternate splicing generate receptors with differing agonist-independent activity," <i>Mol. Pharmacol.</i> , 1996, 49, 422-429	
	DDD	Rakovska, A. et al., "Effect of loxiglumide (CR 1505) on CCK-induced contractions and $^3\text{H}$ -acetylcholine release from guinea-pig gallbladder," <i>Neuropeptides</i> , 1993, 25(5), 271-276	
	DDE	Ren, Q. et al., "Constitutive active mutants of the $\alpha_2$ -adrenergic receptor," <i>J. Biol. Chem.</i> , 1993, 268, 16483-16487	
	DDF	Reynolds, E.E. et al., "Pharmacological characterization of PD 156707, an orally active $\text{ET}_A$ receptor antagonist," <i>J. Pharmacol. Exp. Ther.</i> , 1995, 273(3), 1410-1417	
	DDG	Robbins, L.S. et al., "Pigmentation phenotypes of variant extension locus alleles result from point mutations that alter MSH receptor function," <i>Cell</i> , 1993, 72, 827-834	
	DDH	Rong, S. et al., "Met expression and sarcoma tumorigenicity," <i>Cancer</i> , 1993, 53(22), 5355-5360	
EXAMINER		DATE CONSIDERED	

<b>Form PTO-1449 Modified</b>  List of Patent and Publications Cited by Applicant (Use several sheets if necessary)  U.S. Department of Commerce Patent and Trademark Office		Docket No. 15 US10 CON	Serial No. Not Assigned
		Applicant <b>Dominic P. Behan et al.</b>	
		Filing Date Herewith	Group Not Yet Assigned
<b>OTHER DOCUMENTS (Including Author, Title, Date, Pertinent Pages, Etc.)</b>			
	DDI	Showers, M.O. et al., "Activation of the erythropoietin receptor by the Friend spleen focus-forming virus gp55 glycoprotein induces constitutive protein tyrosine phosphorylation," <i>Blood</i> , 1992, 80(12), 3070-3078	
	DDJ	Skinner, R.H. et al., "Direct measurement of the binding of RAS to neurofibromin using scintillation proximity assay," <i>Anal. Biochem.</i> , 1994, 223, 259-265	
	DDK	Slamon, D.J. et al., "Human breast cancer: correlation of relapse and survival with amplification of the HER-2/neu oncogene," <i>Science</i> , 1987, 235, 177-181	
	DDL	Slamon, D. et al., "Studies of the HER-2/neu proto-oncogene in human breast and ovarian cancer," <i>Science</i> , 1989, 244, 707-712	
	DDM	Salomon, Y. et al., "A highly sensitive adenylate cyclase assay," <i>Anal. Biochem.</i> , 1974, 58, 541-548	
	DDN	Spiegel, A.M., "Defects in G protein-coupled signal transduction in human disease," <i>Ann. Rev. Physiol.</i> , 1995, 58, 143-170	
	DDO	ter Laak, A. et al., "Modelling and mutation studies on the histamine H <sub>1</sub> -receptor agonist binding site reveal different binding modes for H <sub>1</sub> -agonists: Asp <sup>116</sup> (TM3) has a constitutive role in receptor stimulation," <i>J. Computer-Aided Mol. Design</i> , 1995, 9, 319-330	
	DDP	Tiberi, M. et al., "High agonist-independent activity is a distinguishing feature of the dopamine D1B receptor subtype," <i>J. Biol. Chem.</i> , 1994, 269(45), 27925-27931	
	DDQ	Tsujiyama, T. et al., "Constitutive activation of c-kit in FMA3 murine mastocytoma cells caused by deletion of seven amino acids at the juxtamembrane domain," <i>Blood</i> , 1996, 87(1), 273-283	
	DDR	Wang, Z. et al., "Constitutive $\mu$ opioid receptor activation as a regulatory mechanism underlying narcotic tolerance and dependence," <i>Life Sci.</i> , 1994, 54(20), 339-350	
	DDS	Watowich, S.S. et al., "Homodimerization and constitutive activation of the erythropoietin receptor," <i>Proc. Natl. Acad. Sci USA</i> , 1992, 89, 2140-2144	
<b>EXAMINER</b>		<b>DATE CONSIDERED</b>	

<b>Form PTO-1449 Modified</b>  List of Patent and Publications Cited by Applicant (Use several sheets if necessary)  U.S. Department of Commerce Patent and Trademark Office		Docket No. 5.US10.CON	Serial No. Not Assigned
		Applicant <b>Dominic P. Behan et al.</b>	
		Filing Date Herewith	Group Not Yet Assigned
<b>OTHER DOCUMENTS (Including Author, Title, Date, Pertinent Pages, Etc.)</b>			
	DDT	Weber-Nordt, R.M. et al., "Constitutive activation of STAT proteins in primary lymphoid and myeloid leukemia cells and in Epstein-Barr virus (EBV)-related lymphoma cell lines," <i>Blood</i> , 1996, 88(3), 809-816	
	DDU	Webster, M.K. et al., "Constitutive activation of fibroblast growth factor receptor 3 by the transmembrane point mutation found in achondroplasia," <i>EMBO J.</i> , 1996, 15, 520-527	
	DDV	Xu, Y. et al., "Characterization of epidermal growth factor receptor gene expression in malignant and normal human cell lines," <i>Proc. Natl. Acad. Sci. USA</i> , 1984, 81, 7308-7312	
	DDW	Yamada, K. et al., "Substitution of the insulin receptor transmembrane domain with the <i>c-new/erbB2</i> transmembrane domain constitutively activates the insulin receptor kinas <i>in vitro</i> ," <i>J. Biol. Chem.</i> , 1992, 267(18), 12452-12461	
	DDX	Zhang, S. et al., "Identification of Dynorphins as Endogenous Ligands for an Opioid Receptor-Like Orphan Receptor," <i>J. Biol. Chem.</i> , 1995, 270, 22772-22776	
	DEY	Zhen, Z. et al., "Structural and functional domains critical for constitutive activation of the HGF-receptor ( <i>Met</i> )," <i>Oncogene</i> , 1994, 9, 1691-1697	
	DDZ	Gantz, I. et al., "Molecular Cloning of a Novel Melanocortin Receptor," <i>J. Biol. Chem.</i> , 1993, 268(11), 8246-8250	
	DEA	Heiber, M. et al., "Isolation of Three Novel Human Genes Encoding G Protein-Coupled Receptors," <i>DNA and Cell Biology</i> , 1995, 14(1), 25-35	
	DEB	Howard, A.D. et al., "A Receptor in Pituitary and Hypothalamus That Functions in Growth Hormone Release," <i>Science</i> , 1996, 273, 974-977	
	DEC	Iismaa, T.P. et al., "Isolation and Chromosomal Localization of a Novel Human G-Protein-Coupled Receptor (GPR3) Expressed Predominantly in the Central Nervous System," <i>Genomics</i> , 1994, 24, 391-394	
	DED	Itoh, H. et al., "Molecular cloning and sequence determination of cDNAs for $\alpha$ subunits of the guanine nucleotide-binding proteins $G_s$ , $G_i$ , and $G_o$ from rat brain," <i>Proc. Natl. Acad. Sci. USA</i> , 1986, 83, 3776-3780	
<b>EXAMINER</b>		<b>DATE CONSIDERED</b>	

<b>Form PTO-1449 Modified</b>  List of Patent and Publications Cited by Applicant (Use several sheets if necessary)  U.S. Department of Commerce Patent and Trademark Office				Docket No. 5US.10.CON		Serial No. Not Assigned	
				Applicant Dominic P. Behan et al.			
				Filing Date Herewith		Group Not Yet Assigned	
<b>U. S. PATENT DOCUMENTS</b>							
Examiner Initial		Document No.	Date	Name	Class	Subclass	
	EA	5,514,578	05/07/96	Hogness et al.	435	240.2	
	EB	5,532,157	07/02/96	Fink	435	240.2	
	EC	5,573,944	11/12/96	Kirschner et al.	435	252.3	
	ED	5,639,616	06/17/97	Liao et al.	435	7.1	
	EE	5,750,353	05/12/98	Kopin et al.	435	7.21	
<b>FOREIGN PATENT DOCUMENTS</b>							
Examiner Initial		Document No.	Date	Country	Translation YES NO		
	FA	WO 97/11159	09/20/96	PCT	X		
<b>EXAMINER</b>				<b>DATE CONSIDERED</b>			